=> fil hcap FILE 'HCAPLUS' ENTERED AT 10:45:16 ON 30 APR 2007 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2007 AMERICAN CHEMICAL SOCIETY (ACS)

Copyright of the articles to which records in this database refer is held by the publishers listed in the PUBLISHER (PB) field (available for records published or updated in Chemical Abstracts after December 26, 1996), unless otherwise indicated in the original publications. The CA Lexicon is the copyrighted intellectual property of the the American Chemical Society and is provided to assist you in searching databases on STN. Any dissemination, distribution, copying, or storing of this information, without the prior written consent of CAS, is strictly prohibited.

FILE COVERS 1907 - 30 Apr 2007 VOL 146 ISS 19 FILE LAST UPDATED: 29 Apr 2007 (20070429/ED)

STR

New CAS Information Use Policies, enter HELP USAGETERMS for details.

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> d que 16 L3

VAR G1=14/19
VAR G2=H/ME/I-PR
NODE ATTRIBUTES:
CONNECT IS E1 RC AT 12
CONNECT IS E1 RC AT 15
DEFAULT MLEVEL IS ATOM
DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES: RING(S) ARE ISOLATED OR EMBEDDED NUMBER OF NODES IS 21

STEREO ATTRIBUTES: NONE

L5 8 SEA FILE=REGISTRY SSS FUL L3

L6 4 SEA FILE=HCAPLUS ABB=ON PLU=ON L5

=> d 16 ibib abs hitstr tot

L6 ANSWER 1 OF 4 HCAPLUS COPYRIGHT 2007 ACS on STN ACCESSION NUMBER: 2005:397579 HCAPLUS Full-text

DOCUMENT NUMBER:

143:419154

TITLE:

Chemical studies of fish bioluminescence

AUTHOR(S):

Kakoi, Hisae; Okada, Kunisuke

CORPORATE SOURCE:

Faculty of Pharmacy, Meijo University, Tempaku-ku,

Nagoya, 468-8503, Japan

SOURCE:

ITE Letters on Batteries, New Technologies & Medicine

(2005), 6(1), 38-45

CODEN: ILBMF9; ISSN: 1531-2046

PUBLISHER: ITE Inc.

DOCUMENT TYPE: Journal

LANGUAGE: English

Watasenia preluciferin (I), first isolated from the squid Watasenia scintillans, is a compound that plays a key role in the light emitting process of various bioluminescent marine organisms such as squids, shrimps, coelenterates, and fish. In the case of luminous fish, a well-known species is Myctophiformes and Stomiiformes especially the deep-sea photophorespossessing Myctophiformes fish (lantern fish), which is one of the most common and widely distributed luminous fish living in Suruga Bay and all throughout the Sea of Enshu and Kumano. Compound I was isolated either from the liver of Neoscopelus microchir (in Japanese, Sango-iwashi) or from a pair of large nasal photophores from Diaphus gigas (in Japanese, Suito-hadaka) while it was found neither in the photophores of N. microchir nor in the liver of D. gigas. On the other hand, a luciferase active toward Oplophorus luciferin (=Watasenia preluciferin) I was extracted from the flesh of D. gigas, whereas no luciferase active toward I or Cypridina luciferin was found in N. microchir. Later, a new type of bound form of I was isolated from the liver of D. gigas and the structure was established as Diaphus luciferyl β -glucopyranosiduronic acid (II) on the basis of the spectral data and chemical evidence, and by synthesis starting from I. This compound II was also detected in the liver of Diaphus watasei (in Japanese, Hadaka-iwashi) and other examined Myctophiformes fish, but not in the liver of N. microchir. It is uncertain as to which system is more favorable for the fish bioluminescence, however, as far as I is concerned, the Diaphus bioluminescent system is comparable to that of Watasenia or Oplophorus, and not to that of Cypridina as previously observed by Tsuji et al. in 1971.

IT 65417-16-5P

CN

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(intermediate in preparation of luciferyl β -glucopyranosiduronic acid)

RN 65417-16-5 HCAPLUS

Imidazo[1,2-a]pyrazin-3-ol, 6-[4-(acetyloxy)phenyl]-2-[[4(acetyloxy)phenyl]methyl]-8-(phenylmethyl)-, acetate (ester) (9CI) (CA
INDEX NAME)

REFERENCE COUNT:

THERE ARE 18 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L6 ANSWER 2 OF 4 HCAPLUS COPYRIGHT 2007 ACS on STN ACCESSION NUMBER: 2003:376823 HCAPLUS Full-text

DOCUMENT NUMBER:

138:365147

TITLE:

Compositions, methods and kits pertaining to

luminescent compounds

INVENTOR(S):

Wood, Keith; Hawkins, Erika; Scurria, Mike; Klaubert,

Dieter

PATENT ASSIGNEE(S):

Promega Corporation, USA

SOURCE:

PCT Int. Appl., 60 pp.

DOCUMENT TYPE:

CODEN: PIXXD2

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

	PAT	CENT I	NO.			KIN	D	DATE			APPL	ICAT	ION 1	NO.		D.	ATE	
	WO	2003	0401	00		A1	_	2003	0515	1	WO 2	002-	US34:	972		2	0021	101
		W:	ΑE,	AG,	AL,	AM,	AT,	ΑU,	ΑZ,	BA,	BB,	BG,	BR,	BY,	ΒZ,	CA,	CH,	CN,
			CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	ES,	FI,	GB,	GD,	GE,	GH,
			GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	KP,	KR,	KZ,	LC,	LK,	LR,
			LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NO,	NZ,	OM,	PH,
			PL,	PT,	RO,	RU,	SD,	SE,	SG,	SI,	SK,	SL,	TJ,	TM,	TN,	TR,	TT,	TZ,
			UA,	UG,	UZ,	VC,	VN,	YU,	ZA,	ZM,	zw							
		RW:	GH,	GM,	KE,	LS,	MW,	MZ,	SD,	SL,	SZ,	TZ,	UG,	ZM,	ZW,	ΑT,	BE,	BG,
								EE,										
			PT,	SE,	SK,	TR,	BF,	ВJ,	CF,	CG,	CI,	CM,	GA,	GN,	GQ,	GW,	ML,	MR,
			NE,	SN,	TD,	TG												
	US	2003	1530	90		A1		2003	0814	1	US 2	001-	5348	2		2	0011	102
	CA	2462	506			A1		2003	0515	(CA 2	002-	2462	506		2	0021	101
	AU	2002	3634	24		A1		2003	0519		AU 2	002-	3634	24		2	0021	101
	EP	1451	155			A1		2004	0901	1	EP 2	002-	8028	15		2	0021	101
		R:	AT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR,	IT,	LI,	LU,	NL,	SE,	MC,	PT,
			IE,	SI,	LT,	LV,	FI,	RO,	MK,	CY,	AL,	TR,	BG,	CZ,	EE,	SK		
	CN	1612	860			Α		2005	0504	(CN 2	002-	8266'	77		2	0021	101
	JP	2005	5159	77		T		2005	0602	,	JP 2	003-	5421	46		2	0021	101
PRIO	RIT	APP	LN.	INFO	.:					1	US 2	001-	53482	2	7	A 2	0011	102
										1	WO 2	002-1	US349	972	7	w 2	0021	101

OTHER SOURCE(S): MARPAT 138:365147

A method of measuring the enzymic activity of a luciferase includes contacting a luminogenic protein, such as a luciferase, with a protected luminophore to form a composition; and detecting light produced from the composition The protected luminophore provides increased stability and improved signal-tobackground ratios relative to the corresponding unmodified coelenterazine.

IT 65417-16-5P 524066-91-9P 524066-92-0P 524066-93-1P 524066-94-2P 524066-95-3P

524066-96-4P

RL: ARU (Analytical role, unclassified); SPN (Synthetic preparation); ANST (Analytical study); PREP (Preparation)

(compns., methods and kits pertaining to luminescent compds.)

RN 65417-16-5 HCAPLUS

CN Imidazo[1,2-a]pyrazin-3-ol, 6-[4-(acetyloxy)phenyl]-2-[[4-

(acetyloxy)phenyl]methyl]-8-(phenylmethyl)-, acetate (ester) (9CI) INDEX NAME)

RN 524066-91-9 HCAPLUS

CN Imidazo[1,2-a]pyrazin-3-ol, 6-[4-(acetyloxy)phenyl]-2,8-bis(phenylmethyl)-, acetate (ester) (9CI) (CA INDEX NAME)

RN 524066-92-0 HCAPLUS

CN Phenol, 4-[3-[(acetyloxy)methoxy]-2-[[4-(acetyloxy)phenyl]methyl]-8-(phenylmethyl)imidazo[1,2-a]pyrazin-6-yl]-, acetate (ester) (9CI) (CA INDEX NAME)

RN 524066-93-1 HCAPLUS

CN Butanoic acid, 4-[3-[(1-oxobutoxy)methoxy]-2-[[4-(1-oxobutoxy)phenyl]methyl]-8-(phenylmethyl)imidazo[1,2-a]pyrazin-6-yl]phenyl ester (9CI) (CA INDEX NAME)

$$n-Pr$$
 CH_2
 CH_2
 Ph
 CH_2

RN 524066-94-2 HCAPLUS

CN Phenol, 4-[3-[(acetyloxy)methoxy]-2,8-bis(phenylmethyl)imidazo[1,2-a]pyrazin-6-yl]-, acetate (ester) (9CI) (CA INDEX NAME)

RN 524066-95-3 HCAPLUS

Propanoic acid, 2,2-dimethyl-, [[6-(4-hydroxyphenyl)-2,8-CN bis(phenylmethyl)imidazo[1,2-a]pyrazin-3-yl]oxy]methyl ester (9CI) (CA INDEX NAME)

524066-96-4 HCAPLUS RN

CN Methanol, [[6-phenyl-2,8-bis(phenylmethyl)imidazo[1,2-a]pyrazin-3-yl]oxy]-, acetate (ester) (9CI) (CA INDEX NAME)

REFERENCE COUNT: 8 THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L6 HCAPLUS COPYRIGHT 2007 ACS on STN ANSWER 3 OF 4

ACCESSION NUMBER:

2003:108790 HCAPLUS Full-text

DOCUMENT NUMBER:

139:129758

TITLE:

Coelenterazine derivatives for improved solution

solubility

AUTHOR(S):

Hawkins, Erika M.; O'Grady, Michael; Klaubert, Dieter;

Scurria, Michael; Good, Troy; Stratford, Cathy;

Flemming, Rod; Simpson, Dan; Wood, Keith V.

CORPORATE SOURCE:

Promega Corporation, Madison, WI, 53715, USA

SOURCE:

Bioluminescence & Chemiluminescence: Progress &

Current Applications, [Proceedings of the Symposium on

Bioluminescence and Chemiluminescence], 12th, Cambridge, United Kingdom, Apr. 5-9, 2002 (2002), 149-152. Editor(s): Stanley, Philip E.; Kricka, Larry

J. World Scientific Publishing Co. Pte. Ltd.:

Singapore, Singapore.

CODEN: 69DPGZ; ISBN: 981-238-156-2

DOCUMENT TYPE: LANGUAGE:

Conference English

AB Intracellular luminescent techniques requiring coelenterazine, such as bioluminescence resonance energy transfer (BRET), calcium detection, and intracellular reporter measurements, must accommodate the poor stability of this substrate in physiol. buffered solns. Coelenterazine degradation leads both to loss of luminescence over time, and increased background luminescence caused by enzyme-independent oxidation (autoluminescence). Both conditions limit luminescence sensitivity by reducing the signal-to-noise ratio. Coelenterazine can be stabilized by derivatizing the enol oxygen with an acyl oxymethyl ether. This prevents spontaneous oxidation in solution while making the substrate available intracellularly upon cleavage of the blocking group by endogenous esterases. We will describe the stability of pivaloyl oxymethyl coelenterazine-h (POM coelenterazine-h), and the effect of POM coelenterazineh on intracellular luminescence, autoluminescence, and luminescent reaction kinetics. Also, we will present the characteristics of two other coelenterazine derivs.

RN 524066-95-3 HCAPLUS

CN Propanoic acid, 2,2-dimethyl-, [[6-(4-hydroxyphenyl)-2,8-bis(phenylmethyl)imidazo[1,2-a]pyrazin-3-yl]oxy]methyl ester (9CI) (CA INDEX NAME)

RN 566945-96-8 HCAPLUS

CN Butanoic acid, [[6-(4-hydroxyphenyl)-2,8-bis(phenylmethyl)imidazo[1,2-a]pyrazin-3-yl]oxy]methyl ester (9CI) (CA INDEX NAME)

L6 ANSWER 4 OF 4 HCAPLUS COPYRIGHT 2007 ACS on STN ACCESSION NUMBER: 1978:50764 HCAPLUS Full-text

DOCUMENT NUMBER:

88:50764

TITLE:

Complete structure of Renilla luciferin and luciferyl

sulfate

AUTHOR(S):

Inoue, Shoji; Kakoi, Hisae; Murata, Mikiko; Goto,

Toshio; Shimomura, Osamu

CORPORATE SOURCE:

SOURCE:

Fac. Pharm., Meijo Univ., Nagoya, Japan Tetrahedron Letters (1977), (31), 2685-8

CODEN: TELEAY; ISSN: 0040-4039

DOCUMENT TYPE:

Journal

LANGUAGE:

English

GI

AB Examination of Renilla exts. showed that Renilla luciferin is coëlenterazine (I). The structure of natural luciferyl sulfate was determined as II by comparison of natural and synthetic II. II was synthesized from I by sequential treatment with (AcO)2O, MeOH/NH3, and pyridine-SO3 complex and hydrolysis with MeOH/NaOH.

IT 65417-16-5P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(preparation and hydrolysis of)

RN 65417-16-5 HCAPLUS

CN Imidazo[1,2-a]pyrazin-3-ol, 6-[4-(acetyloxy)phenyl]-2-[[4-(acetyloxy)phenyl]methyl]-8-(phenylmethyl)-, acetate (ester) (9CI) (CA INDEX NAME)

=> fil marpat

FILE 'MARPAT' ENTERED AT 10:45:32 ON 30 APR 2007
USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.
PLEASE SEE "HELP USAGETERMS" FOR DETAILS.
COPYRIGHT (C) 2007 American Chemical Society (ACS)

FILE CONTENT: 1961-PRESENT VOL 146 ISS 18 (20070427/ED)

SOME MARPAT RECORDS ARE DERIVED FROM INPI DATA FOR 1961-1987

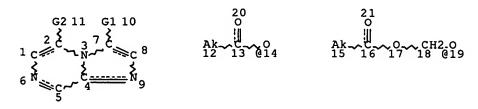
MOST RECENT CITATIONS FOR PATENTS FROM MAJOR ISSUING AGENCIES (COVERAGE TO THESE DATES IS NOT COMPLETE):

US 2007060644 15 MAR 2007
DE 102006023116 15 MAR 2007
EP 1762248 14 MAR 2007
JP 2007059877 08 MAR 2007
WO 2007030662 15 MAR 2007
GB 2429975 14 MAR 2007
FR 2890657 16 MAR 2007
RU 2295953 27 MAR 2007
CA 2556850 24 FEB 2007

Expanded G-group definition display now available.

=> d que 116

L3 STR



VAR G1=14/19
VAR G2=H/ME/I-PR
NODE ATTRIBUTES:
CONNECT IS E1 RC AT 12
CONNECT IS E1 RC AT 15
DEFAULT MLEVEL IS ATOM
DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:

RING(S) ARE ISOLATED OR EMBEDDED

NUMBER OF NODES IS 21

STEREO ATTRIBUTES: NONE

L5 8 SEA FILE=REGISTRY SSS FUL L3

L6 4 SEA FILE=HCAPLUS ABB=ON PLU=ON L5

L15 2 SEA FILE=MARPAT SSS FUL L3

L16 1 SEA FILE=MARPAT ABB=ON PLU=ON L15 NOT L6

=> d l16 ibib abs qhit tot

L16 ANSWER 1 OF 1 MARPAT COPYRIGHT 2007 ACS on STN 135:371764 MARPAT Full-text ACCESSION NUMBER:

TITLE: Preparation of aminopyrazines and imidazolopyrazinones as

INVENTOR(S): Marchand-Brynaert, Jacqueline; Cavalier,

Jean-Francois; Rees, Jean-Francois; De Tollenaere,

Catherine; Burton, Maggi

PATENT ASSIGNEE(S): Universite Catholique de Louvain, Belg.

SOURCE: PCT Int. Appl., 57 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent. LANGUAGE: English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PA'	TENT	NO.		KI	ND :	DATE		•	A.	PPLI	CATI	ON NO	٥.	DATE			
WO	2001	0878	53	A	1	2001	1122		W	20	01-E	P558	3	2001	0516		
	W:	ΑE,	AG,	AL,	AM,	AT,	AU,	AZ,	BA,	BB,	BG,	BR,	BY,	BZ,	CA,	CH,	CN,
		CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EE,	ES,	FI,	GB,	GD,	GE,	GH,	GM,	HR,
		HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	KP,	KR,	ΚZ,	LC,	LK,	LR,	LS,	LT,
		LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NO,	NZ,	PL,	PT,	RO,	RU,
		SD,	SE,	SG,	SI,	SK,	SL,	TJ,	TM,	TR,	TT,	ΤZ,	UA,	ŪG,	US,	UZ,	VN,
		YU,	ZA,	ZW,	AM,	AZ,	BY,	KG,	ΚZ,	MD,	RU,	TJ,	TM				
	RW:	GH,	GM,	KE,	LS,	MW,	MZ,	SD,	SL,	SZ,	TZ,	UG,	ZW,	AT,	BE,	CH,	CY,
		DE,	DK,	ES,	FI,	FR,	GB,	GR,	ΙE,	IT,	LU,	MC,	NL,	PT,	SE,	TR,	BF,
														TD,			
EP	1292	580		Α	1 :	2003	0319		E	P 20	01-9	43383	3	2001	0516		
	R:	AT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR,	IT,	LI,	LU,	NL,	SE,	MC,	PT,
		IE,	SI,	LT,	LV,	FI,	RO,	MK,	CY,	AL,	TR	•					-
US	2004	0342	25	A	1 :	2004	0219		U:	5 20	03-2	76391	3	2003	0728		
PRIORIT	Y APP	LN.	INFO	.:					E	P 20	00-8	7010	7	2000	0517		
									E	P 20	00-8	7029:	3	2000	1212		
									W	20	01-E	P5588	3	2001	0516		

CASREACT 135:371764 OTHER SOURCE(S):

Antioxidants, 5 2-amino-(p-hydroxyphenyl)pyrazines and 3 (p-hydroxyphenyl)-3,7-dihydroimidazo[1,2-a]pyrazin-3-ones were prepared and claimed useful in diagnostic procedures, as food additives, polymer additives and as UV screens in cosmetics. E.g., 2-amino-3,5- dibromopyrazine was treated with pmethoxyphenylboronic acid in the presence of bis(benzonitrile)palladium dichloride and 1,4- bis(diphenylphosphino) butane in a solvent mix of EtOH, aqueous sodium carbonate and toluene to give 66% 2-amino-3,5-bis(pmethoxyphenyl)pyrazine, which was demethylated with EtSNa in DMF to give 88% 2-amino-3,5-bis(p-hydroxyphenyl)pyrazine (I). In tests on inhibition of lipid peroxidn. 2-aminopyrazines possessing 2 aryl substituents, one of them being a p-hydroxyphenyl in o- or p- position with respect to the amino group, are endowed with antioxidative properties. However, the p-hydroxyphenyl conferred more activity when located at position 5 than at position 3. The presence of p-hydroxyphenyl groups at both positions 3 and 5 as in I produced a very active compound Analogs lacking the free phenol groups showed reduced activities. Corresponding imidazolopyrazinones combined the properties of both the imidazolopyrazinones (delay of the onset of peroxidn.) and the aminopyrazines (lower rate of oxidation after onset).

G6 = OCOMe

Patent location:

claim 1 Note: or prodrugs, or pharmaceutically acceptable

addition salts, or tautomerically isomeric forms

Stereochemistry: or stereochemically isomeric forms

MSTR 3

= OCOMe

Patent location:

Note:

claim 27

or prodrugs, or pharmaceutically acceptable

addition salts, or tautomerically isomeric forms

or stereochemically isomeric forms

Stereochemistry: REFERENCE COUNT:

10 THERE ARE 10 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

INVENTOR NAME SEARCH

=> fil hcap medline embase biosis dissabs wpix FILE 'HCAPLUS' ENTERED AT 10:46:23 ON 30 APR 2007 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2007 AMERICAN CHEMICAL SOCIETY (ACS)

FILE 'MEDLINE' ENTERED AT 10:46:23 ON 30 APR 2007

FILE 'EMBASE' ENTERED AT 10:46:23 ON 30 APR 2007 Copyright (c) 2007 Elsevier B.V. All rights reserved.

FILE 'BIOSIS' ENTERED AT 10:46:23 ON 30 APR 2007 Copyright (c) 2007 The Thomson Corporation

FILE 'DISSABS' ENTERED AT 10:46:23 ON 30 APR 2007

COPYRIGHT (C) 2007 ProQuest Information and Learning Company; All Rights Reserved.

FILE 'WPIX' ENTERED AT 10:46:23 ON 30 APR 2007 COPYRIGHT (C) 2007 THE THOMSON CORPORATION

=> d que 2610 SEA WOOD K/AU OR WOOD K ?/AU OR WOOD KEITH?/AU L17 L18 948 SEA HAWKINS E/AU OR HAWKINS E?/AU OR HAWKINS ERI!A?/AU L19 34 SEA ("SCURRIA M"/AU OR "SCURRIA M A"/AU OR "SCURRIA M S"/AU OR "SCURRIA MICHAEL"/AU OR "SCURRIA MICHAEL A"/AU OR "SCURRIA MIKE"/AU) L20 198 SEA ("KLAUBERT D"/AU OR "KLAUBERT D H"/AU OR "KLAUBERT D K"/AU OR "KLAUBERT DIETER"/AU OR "KLAUBERT DIETER H"/AU OR "KLAUBERT DIETER HEINZ"/AU) L21 44 SEA (L17 AND (L18 OR L19 OR L20)) OR (L18 AND (L19 OR L20)) OR (L19 AND L20)

=> dup rem 121

PROCESSING COMPLETED FOR L21

L22 24 DUP REM L21 (20 DUPLICATES REMOVED) ANSWERS '1-15' FROM FILE HCAPLUS ANSWERS '16-23' FROM FILE BIOSIS

ANSWER '24' FROM FILE WPIX

=> d 122 ibib ab tot

L22 ANSWER 1 OF 24 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 1

2006:1279865 HCAPLUS Full-text ACCESSION NUMBER:

DOCUMENT NUMBER: 146:57589

TITLE: Luminogenic and fluorogenic compounds and methods to

detect molecules or conditions

INVENTOR(S): Daily, William; Hawkins, Erika;

> Klaubert, Dieter; Liu, Jianquan; Meisenheimer, Poncho; Scurria, Michael; Shultz, John W.; Unch, James; Wood, Keith V.; Zhou, Wenhui;

Valley, Michael P.; Cali, James J.

PATENT ASSIGNEE(S): Promega Corporation, USA SOURCE:

PCT Int. Appl., 328pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English PATENT INFORMATION:

PA'	PATENT NO.					D	DATE			APP	LICAT	ION 1	NO.			ATE	
	2006 2006								1	WO	2006-	US20	731	-		0060	
	W:								BA,	вв	, BG,	BR,	BW,	BY,	BZ,	CA,	ÇH,
											, EC,						
		GE,	GH,	GM,	HR,	HU,	ID,	IL,	IN,	IS	, JP,	KE,	KG,	KM,	KN,	KP,	KR,
		KZ,	LC,	LK,	LR,	LS,	LT,	LU,	LV,	LY	, MA,	MD,	MG,	MK,	MN,	MW,	MX,
		MZ,	NA,	NG,	NI,	NO,	NZ,	OM,	PG,	PH	, PL,	PT,	RO,	RU,	SC,	SD,	SE,
		SG,	SK,	SL,	SM,	SY,	ТJ,	TM,	TN,	TR	, TT,	TZ,	UA,	UG,	US,	UZ,	VC,
		VN,	YU,	ZA,	ZM,	ZW											
	RW:	AT,	BE,	BG,	CH,	CY,	CZ,	DE,	DK,	EE	; ES,	FI,	FR,	GB,	GR,	HU,	ΙE,
		IS,	IT,	LT,	LU,	LV,	MC,	NL,	PL,	PT	, RO,	SE,	SI,	SK,	TR,	BF,	ВJ,
		CF,	CG,	CI,	CM,	GA,	GN,	GQ,	GW,	ML	, MR,	NE,	SN,	TD,	TG,	BW,	GH,
		GM,	KE,	LS,	MW,	MZ,	NA,	SD,	SL,	SZ	, TZ,	UG,	ZM,	ZW,	AM,	AZ,	BY,
		KG,	KZ,	MD,	RU,	TJ,	TM										
US	2007	0157	90		A1		2007	0118	1	US	2006-	4441	45		2	0060	531
PRIORIT	Y APP	LN.	INFO	.:					1	US	2005-	6859	57P	1	P 2	0050	531
									1	US	2005-	6930	34P	1	P 2	0050	621
									1	US	2005-	6929	25P]	P 2	0050	622
									1	US	2006-	7904	55P	1	P 20	0060	407

OTHER SOURCE(S): MARPAT 146:57589

AB A method to detect the presence or amount of at least one mol. in a sample which employs a derivative of luciferin or a derivative of a fluorophore is provided.

L22 ANSWER 2 OF 24 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 2

ACCESSION NUMBER: 2006:1161664 HCAPLUS Full-text

DOCUMENT NUMBER: 146:116710

TITLE: Electrophilic aromatic substituted luciferins as

bioluminescent probes for glutathione S-transferase

assays

AUTHOR(S): Zhou, Wenhui; Shultz, John W.; Murphy, Nancy;

Hawkins, Erika M.; Bernad, Laurent; Good, Troy; Moothart, Leonard; Frackman, Susan; Klaubert, Dieter H.; Bulleit, Robert F.;

Wood, Keith V.

CORPORATE SOURCE: Promega Biosciences Inc., San Luis Obispo, CA, 93401,

USA

SOURCE: Chemical Communications (Cambridge, United Kingdom)

(2006), (44), 4620-4622

CODEN: CHCOFS; ISSN: 1359-7345

PUBLISHER: Royal Society of Chemistry

DOCUMENT TYPE: Journal LANGUAGE: English

AB New highly sensitive latent bioluminescent luciferin substrates were designed and synthesized for monitoring mammalian glutathione S-transferase (GST) and

Schistosoma japonicum enzyme activities.

REFERENCE COUNT: 26 THERE ARE 26 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L22 ANSWER 3 OF 24 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 3

ACCESSION NUMBER: 2006:137803 HCAPLUS Full-text

DOCUMENT NUMBER: 144:384691

TITLE: New Bioluminogenic Substrates for Monoamine Oxidase

Assays

10/053,482 April 30, 2007

Zhou, Wenhui; Valley, Michael P.; Shultz, John; AUTHOR(S):

> Hawkins, Erika M.; Bernad, Laurent; Good, Troy; Good, Dave; Riss, Terry L.; Klaubert,

Dieter H.; Wood, Keith V.

CORPORATE SOURCE: Promega Biosciences Inc., San Luis Obispo, CA, 93401,

SOURCE: Journal of the American Chemical Society (2006),

128(10), 3122-3123

CODEN: JACSAT; ISSN: 0002-7863

PUBLISHER: American Chemical Society

DOCUMENT TYPE: Journal LANGUAGE: English

Novel bioluminogenic substrates were designed for probing monoamine oxidase (MAO) activity based on a simple and effective β -elimination strategy. modifying the amino group and the central core of luciferin derivs., we have developed a series of substrates useful for assays of MAO A or B, or both. One of these substrates, exhibiting low Km values and high signal-tobackground ratios with both isoenzymes, was shown to accurately measure the Ki values of known MAO inhibitors. This substrate is a key component in the development of a highly sensitive homogeneous MAO assay for high-throughput screening (HTS) of compds. in drug discovery and for monitoring MAO activity in complex biol. systems. This design strategy should be applicable to fluorogenic MAO substrates and could broaden the structural requirements of substrates for other enzyme assays.

REFERENCE COUNT: 15 THERE ARE 15 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L22 ANSWER 4 OF 24 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 4

2006:1268397 HCAPLUS Full-text ACCESSION NUMBER:

DOCUMENT NUMBER: 146:200635

TITLE: A bioluminescent assay for monoamine oxidase activity

AUTHOR(S): Valley, Michael P.; Zhou, Wenhui; Hawkins, Erika

M.; Shultz, John; Cali, James J.; Worzella,

Tracy; Bernad, Laurent; Good, Troy; Good, Dave; Riss,

Terry L.; Klaubert, Dieter H.; Wood,

Keith V.

CORPORATE SOURCE: Promega Corporation, Madison, WI, 53711, USA

SOURCE: Analytical Biochemistry (2006), 359(2), 238-246

CODEN: ANBCA2; ISSN: 0003-2697

PUBLISHER: Elsevier DOCUMENT TYPE: Journal LANGUAGE: English

This article describes a novel two-step homogeneous bioluminescent assay for AB monoamine oxidase (MAO) that is simple, sensitive, and amenable to highthroughput screening. In the first step, MAO reacts with an aminopropylether analog of Me ester luciferin. In the second step, a luciferin detection reagent inactivates MAO and converts the product of the first step into a luminescent signal. The amount of light produced is proportional to the amount of MAO and the time of incubation in the first step, but the luminescent signal is stable in the second step with a half-life greater than 5 h. The assay has high precision, is more sensitive than current fluorescent methods, and can accurately measure the binding consts. of known substrates and inhibitors. An automated screen of the Sigma-RBI Library of Pharmacol. Active Compds. (LOPAC1280) revealed a surprisingly high percentage of MAO inhibitors (16%) with a low false hit rate (0.9%). This implies that a significant number of compds. interact with the MAO enzymes and suggests that it is important to include MAO assays in drug metabolism studies. Other advantages of this bioluminescent assay over comparable fluorescent assays are discussed.

REFERENCE COUNT: 29 THERE ARE 29 CITED REFERENCES AVAILABLE FOR THIS

RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L22 ANSWER 5 OF 24 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 5

ACCESSION NUMBER: 2005:1292638 HCAPLUS Full-text

DOCUMENT NUMBER: 144:33522

TITLE: Substrate-binding, catalytically inactive hydrolases

as carriers for the immobilization of fusion proteins

INVENTOR(S): Darzins, Aldis; Encell, Lance; Johnson, Tonny;

Klaubert, Dieter; Los, Georgyi V.; Mcdougall,

Mark; Wood, Keith V.; Wood, Monika G.;

Zimprich, Chad

PATENT ASSIGNEE(S): USA

U.S. Pat. Appl. Publ., 121 pp., Cont.-in-part of U.S. SOURCE:

Ser. No. 768,976.

CODEN: USXXCO

DOCUMENT TYPE:

Patent

LANGUAGE: English FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT	NO.		KIN	D	DATE			APPL	ICAT				D.	ATE	
US 2005	272114	-	A1	_	2005	1208		US 2					2	 0041	 206
US 2004	214258		A1		2004	1028		US 2	004-	7689	76		2	0040	130
US 2006	024808		A1		2006			US 2	005-	1941	10		_	0050	
CA 2575	611		A1		2006	0908		CA 2						0050	
WO 2006	093529		A2		2006	0908		WO 2					_	0050	
WO 2006					2007						•••		_	0000	, 23
W:		AL.					BA.	BB.	BG.	BR.	BW.	BY.	B7.	CA.	CH.
	CN, CO,														
	GE, GH,														
	LC, LK,											-	-	-	-
	NG, NI,														
	SL, SM,														
	ZA, ZM,		10,	111,	111,	111,	11,	14,	UA,	00,	05,	02,	٧٠,	AIA,	10,
DW•	AT, BE,		CH	CV	C7	חבי	אמ		FC	ET	מים	CP	CD	1111	TE
1744 .	IS, IT,														
	CF, CG,														
	GM, KE,					SU,	эь,	54,	TZ,	UG,	ZM,	ZW,	AM,	AZ,	BY,
110 2007	KG, KZ,	•	•	•		0410			006	- 003	۰.		_	0060	
	087400		ΑŢ		2007	0419		US 2						0060	
PRIORITY APP	LN. INFO).:						US 2						0030	
								US 2						0030	
								US 2			. •	_	_	0040	
								US 2		_				0040	
								US 2				_	_	0041	
סדעבם פסווסכב					1 / / .			WO 2	005-1	JS27	307	1	W 2	0050	729

OTHER SOURCE(S): MARPAT 144:33522

Hydrolase variants that retain substrate binding, and capable of forming a covalent bond with a substrate, but lacking the catalytic activity to release the hydrolysis products are described for use in the immobilization of proteins onto surfaces carrying a substrate for the hydrolase are described. The binding of the hydrolase to substrate is more stable than that of the wild type enzyme. The catalytically inactive variant has at least two amino acid substitutions. Substrates for hydrolases comprising one or more functional groups are also provided, as well as methods of using the mutant hydrolase and the substrates of the invention. Also provided is a fusion protein capable of forming a stable bond with a substrate and cells which express the fusion protein. Development of a catalytically inactive variant of the haloalkane dehalogenase of Rhodococcus rhodochrous is demonstrated. Use of fusion

products with fluorescent proteins and enzymes in imaging in vivo are demonstrated.

L22 ANSWER 6 OF 24 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 6

ACCESSION NUMBER:

2005:291435 HCAPLUS Full-text

DOCUMENT NUMBER:

143:341532

TITLE:

Homogeneous, bioluminescent protease assays: Caspase-3

as a model

AUTHOR(S):

O'Brien, Martha A.; Daily, William J.; Hesselberth, P.

Eric; Moravec, Richard A.; Scurria, Michael A. ; Klaubert, Dieter H.; Bulleit, Robert F.;

Wood, Keith V.

CORPORATE SOURCE:

Promega Corporation, Madison, WI, USA

SOURCE:

Journal of Biomolecular Screening (2005), 10(2),

137-148

CODEN: JBISF3; ISSN: 1087-0571

PUBLISHER:

Sage Publications

DOCUMENT TYPE:

Journal

LANGUAGE: English

Using caspase-3 as a model, the authors have developed a strategy for highly AB sensitive, homogeneous protease assays suitable for high-throughput, automated applications. The assay uses peptide-conjugated aminoluciferin as the protease substrate and a firefly luciferase that has been molecularly evolved for increased stability. By combining the proluminescent caspase-3 substrate, Z-DEVD-aminoluciferin, with a stabilized luciferase in a homogeneous format, the authors developed an assay that is significantly faster and more sensitive than fluorescent caspase-3 assays. The assay has a single-step format, in which protease cleavage of the substrate and luciferase oxidation of the aminoluciferin occurs simultaneously. Because these processes are coupled, they rapidly achieve steady state to maintain stable luminescence for several hours. Maximum sensitivity is attained when this steady state occurs; consequently, this coupled-enzyme system results in a very rapid assay. homogeneous format inherently removes trace contamination by free aminoluciferin, resulting in extremely low background and yielding exceptionally high signal-to-noise ratios and excellent Z' factors. Another advantage of a luminescent format is that it avoids problems of cell autofluorescence or fluorescence interference that can be associated with synthetic chemical and natural product libraries. This bioluminescent, homogeneous format should be widely applicable to other protease assays.

REFERENCE COUNT:

THERE ARE 25 CITED REFERENCES AVAILABLE FOR THIS 25 RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L22 ANSWER 7 OF 24 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 7

ACCESSION NUMBER:

2005:150211 HCAPLUS Full-text

TITLE:

Analytical biotechnology

AUTHOR(S):

Wood, Keith V.; Klaubert, Dieter H.

CORPORATE SOURCE:

Promega Corporation, Madison, WI, 53711, USA

SOURCE:

Current Opinion in Biotechnology (2005), 16(1), 1-2

CODEN: CUOBE3; ISSN: 0958-1669

PUBLISHER:

Elsevier Ltd.

DOCUMENT TYPE:

Journal; Editorial

LANGUAGE:

English

AB Unavailable

L22 ANSWER 8 OF 24 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 8

ACCESSION NUMBER:

2004:698252 HCAPLUS Full-text

DOCUMENT NUMBER:

141:187324

TITLE:

Methods and kits for dual enzymatic assays whereby

10/053,482 April 30, 2007

light is quenched from luminescent reactions

INVENTOR(S): Hawkins, Erika; Butler, Braeden; Wood,

Keith V.

PATENT ASSIGNEE(S): Promega Corporation, USA SOURCE: PCT Int. Appl., 77 pp.

CODEN: PIXXD2

DOCUMENT TYPE: LANGUAGE:

Patent English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

		CENT 1									APPL							
		2004				•											0040	
		W:	ΑE,	AG,	AL,	AM,	AT,	ΑU,	ΑZ,	BA,	BB,	BG,	BR,	BW,	BY,	BZ,	CA,	CH,
			CN,	CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	EG,	ES,	FI,	GB,	GD,
			GE,	GH,	GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	KP,	KR,	KZ,	LC,
									MA,									
		RW:	BW,	GH,	GM,	KE,	LS,	MW,	MZ,	SD,	SL,	SZ,	TZ,	ŪG,	ZM,	ZW,	AT,	BE,
			BG,	CH,	CY,	CZ,	DE,	DK,	EE,	ES,	FI,	FR,	GB,	GR,	HU,	IE,	IT,	LU,
			MC,	NL,	PT,	RO,	SE,	SI,	SK,	TR,	BF,	ВJ,	CF,	CG,	CI,	CM,	GA,	GN,
			GQ,	GW,	ML,	MR,	NE,	SN,	TD,	TG								
	ΑU	2004	2109	82		A1		2004	0826		AU 2	004-	2109	82		2	0040	212
		2515							0826									
	US	2004	2243	77		A1		2004	1111	1	US 2	004-	7774	61		2	0040	212
	ΕP	1592	805			A1		2005	1109		EP 2	004-	7105	94		2	0040	212
		R:	AT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR,	İT,	LI,	LU,	NL,	SE,	MC,	PT,
									MK,									
	JP	2006	5174	13		T		2006	0727		JP 2	006-	5035	13		2	0040	212
PRIOR	(TI	APP:	LN.	INFO	. :					1	US 2	003-	4470	65P	1	P 20	0030	212
										1	WO 2	004-1	US40'	75	7	W 2	0040	212

AB The present invention relates to single and dual reporter luminescence assays utilizing reagents to quench an optical, e.g., an enzyme-mediated luminescence, reaction. In one embodiment of the invention, a reagent is added to an assay which selectively quenches a first enzyme-mediated luminescence reaction without affecting a subsequent distinct enzyme-mediated luminescent reaction(s). An assay kit containing one or more selective quench reagents, and compns. comprising the quench reagent(s), are also provided.

L22 ANSWER 9 OF 24 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 9

ACCESSION NUMBER: 2004:698213 HCAPLUS Full-text

DOCUMENT NUMBER: 141:221282

TITLE: Mutant Rhodococcus dehalogenase and functionalized

chloroalkane substrates useful for covalent tethering

of functional groups to proteins

INVENTOR(S): Wood, Keith V.; Los, Georgyi V.; Bulleit,

Robert F.; Klaubert, Dieter; Mcdougall,

Mark; Zimprich, Chad

PATENT ASSIGNEE(S): Promega Corporation, USA

SOURCE: PCT Int. Appl., 185 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2004072232	A2	20040826	WO 2004-US2607	20040130

```
WO 2004072232
                          A9
                                20041014
     WO 2004072232
                          A3
                                20050127
         W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
             CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,
             GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC,
             LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI
         RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE,
             BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU,
             MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN,
             GQ, GW, ML, MR, NE, SN, TD, TG
    AU 2004211584
                          A1
                                20040826
                                            AU 2004-211584
                                                                    20040130
     CA 2514564
                          A1
                                20050726
                                            CA 2004-2514564
                                                                    20040130
    EP 1594962
                          A2
                                20051116
                                            EP 2004-707032
                                                                    20040130
            AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK
     CN 1764721
                          Α
                                20060426
                                            CN 2004-80008194
                                                                    20040130
PRIORITY APPLN. INFO.:
                                            US 2003-444094P
                                                                 Р
                                                                    20030131
                                            US 2003-474659P
                                                                 Ρ
                                                                    20030530
                                            WO 2004-US2607
                                                                 W
                                                                    20040130
```

OTHER SOURCE(S): MARPAT 141:221282

A mutant hydrolase optionally fused to a protein of interest is provided. Thus, Rhodococcus haloalkane dehalogenase DhaA with His-272 substituted with Phe is capable of forming a bond with a chloroalkane substrate for the corresponding nonmutant (wild-type) hydrolase which is more stable than the bond formed between the wild-type hydrolase and the substrate. The chloroalkane substrate contains a functional group which binds Ca2+ or K+ , or Na+, is pH sensitive, is a radionuclide, is electron opaque, is a chromophore or fluorophore, is a MRI contrast agent, is a substance that fluoresces in the presence of NO, or is sensitive to reactive oxygen. Substrates for hydrolases comprising one or more functional groups are synthesized comprising TAMRA-, FAM-, and ROX.5-C14H24O4-Cl or biotin-C18H32O4-Cl, as methods of using the mutant DhaA and the substrates of the invention for cell imaging in vivo are provided. Mutant Staphylococcus aureus β -lactamase (blaZ)-based tethering of functional groups is also demonstrated. Also provided is a fusion protein capable of forming a stable bond with a substrate and cells which express the fusion protein.

L22 ANSWER 10 OF 24 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 10

2004:570131 HCAPLUS Full-text ACCESSION NUMBER:

DOCUMENT NUMBER:

TITLE: Improving the accuracy of luciferase-based assays for

high throughput screening by using tolerance

enhancement agents

INVENTOR(S): Hawkins, Erika; Cali, James J.; Ho, Samuel

Kin Sang; O'Brien, Martha; Somberg, Richard; Bulleit,

Robert F.; Wood, Keith V.

PATENT ASSIGNEE(S):

Promega Corporation, USA PCT Int. Appl., 68 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

SOURCE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2004059294	A2	20040715	WO 2003-US41454	20031223
WO 2004059294	A3	20060629		
W: AE, AG, AL,	AM, AT	, AU, AZ, BA	BB, BG, BR, BY, BZ,	CA, CH, CN,

```
CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
             GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
             LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM,
             PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN,
             TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
         RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ,
             BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE,
             ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK,
             TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
     CA 2508072
                          A1
                                20040715
                                            CA 2003-2508072
                                                                    20031223
    AU 2003300008
                          A1
                                20040722
                                            AU 2003-300008
                                                                    20031223
     US 2005026171
                          A1
                                20050203
                                            US 2003-746995
                                                                    20031223
     EP 1588143
                          A2
                                20051026
                                            EP 2003-800272
                                                                    20031223
             AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK
     JP 2006517401
                          Т
                                20060727
                                            JP 2005-510073
                                                                    20031223
PRIORITY APPLN. INFO.:
                                            US 2002-436173P
                                                                    20021223
                                                                 Ρ
                                            US 2003-444264P
                                                                 Ρ
                                                                    20030131
                                            US 2003-447334P
                                                                 Ρ
                                                                    20030213
                                            WO 2003-US41454
                                                                 W
                                                                    20031223
```

AB The invention concerns methods and kits for improving the accuracy of luciferase-based assays for high throughput screening of compound libraries by reducing the number of false hits. A method and kit is provided for enhancing the tolerance of an assay reagent to compds. in an assay sample, the assay reagent including a luciferase enzyme. The method includes contacting the luciferase with a tolerance enhancement agent in an amount sufficient to substantially protect luciferase enzyme activity from interference of the compound and minimize interference by at least about 10% relative to an assay not having tolerance enhancement agent. Tolerance-enhancing effect of detergents on the inhibition of luciferase was studied. Minimization of false hit occurrence using tolerance enhancement agents such as detergents was demonstrated.

```
L22 ANSWER 11 OF 24
                     HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 11
```

ACCESSION NUMBER:

2004:270174 HCAPLUS Full-text

DOCUMENT NUMBER:

140:299425

TITLE:

Luminescent cytochrome P 450 assay using luciferase, luciferin derivatives and pyrophosphatase, and drug

screening applications

INVENTOR(S):

Cali, James J.; Klaubert, Dieter; Daily,

William; Ho, Samuel Kin Sang; Frackman, Susan;

Hawkins, Erika; Wood, Keith V.

PATENT ASSIGNEE(S):

Promega Corporation, USA PCT Int. Appl., 130 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

SOURCE:

English

FAMILY ACC. NUM. COUNT:

PATENT NO.			KIN	D	DATE			APPL:	ICAT:	ION I	NO.		D	ATE	
				-			•								
WO 2004027	378		A2		2004	0401	1	WO 2	003-1	US29	078		2	00309	916
WO 2004027	0 2004027378 W: AE, AG, A				2004	1125									
W: AE	, AG,	AL,	AM,	AT,	AU,	AZ,	BA,	BB,	BG,	BR,	BY,	BZ,	CA,	CH,	CN,
CC	, CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	ES,	FI,	GB,	GD,	GE,	GH,
GM	, HR,	HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	KP,	KR,	KZ,	LC,	LK,	LR,
LS	, LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NI,	NO,	NZ,	OM,
PG	, PH,	PL,	PT,	RO,	RU,	SC,	SD,	SE,	SG,	SK,	SL,	SY,	TJ,	TM,	TN,

```
TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY,
             KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES,
             FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR,
             BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
     CA 2497560
                          A1
                                20040401
                                            CA 2003-2497560
                                                                    20030916
    AU 2003267245
                          A1
                                20040408
                                            AU 2003-267245
                                                                    20030916
                                20050629
                          A2
    EP 1546162
                                          EP 2003-749715
                                                                    20030916
             AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK
     JP 2006508339
                          Т
                                20060309
                                            JP 2004-537859
     US 2004171099
                          A1
                                20040902
                                            US 2003-665314
                                                                    20030919
PRIORITY APPLN. INFO .:
                                            US 2002-412254P
                                                                 P
                                                                    20020920
                                            US 2003-483309P
                                                                 Ρ
                                                                    20030627
                                            WO 2003-US29078
                                                                    20030916
```

OTHER SOURCE(S): MARPAT 140:299425

AB The present invention provides methods, compns., substrates, and kits useful for analyzing the metabolic activity in cells, tissue, and animals and for screening test compds. for their effect on cytochrome P 450 activity. particular, a one-step and two-step methods using luminogenic mols., e.g. luciferin or coelenterazines, that are cytochrome P 450 substrates and that are also bioluminescent enzyme, e.g., luciferase, pro-substrates are provided. Upon addition of the luciferin derivative or other luminogenic mol. into a P 450 reaction, the P 450 enzyme metabolizes the mol. into a bioluminescent enzyme substrate, e.g., luciferin and/or luciferin derivative metabolite, in a P 450 reaction. The resulting metabolite(s) serves as a substrate of the bioluminescent enzyme, e.g., luciferase, in a second light-generating reaction. Luminescent cytochrome P 450 assays with low background signals and high sensitivity are disclosed and isoform selectivity is demonstrated. present invention also provides an improved method for performing luciferase reactions which employs added pyrophosphatase to remove inorg. pyrophosphate, a luciferase inhibitor which may be present in the reaction mixture as a contaminant or may be generated during the reaction. The present method further provides a method for stabilizing and prolonging the luminescent signal in a luciferase-based assay using luciferase stabilizing agents such as reversible luciferase inhibitors.

L22 ANSWER 12 OF 24 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 12

ACCESSION NUMBER: 2003:633682 HCAPLUS Full-text

139:193612 DOCUMENT NUMBER:

TITLE: Bioluminescent protease assay using aminoluciferin

linked to peptide substrate and luciferase

INVENTOR(S): O'Brian, Martha; Wood, Keith; Klaubert,

Dieter; Daily, Bill

PATENT ASSIGNEE(S): Promega Corporation, USA SOURCE: PCT Int. Appl., 55 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT:

PAT	ENT	NO.			KIN	D .	DATE			APPL	ICAT	ION !	NO.		Di	ATE	
						-			•								
WO	2003	0666	11		A1		2003	0814	1	WO 2	003-1	US29.	36		20	0030	131
	W:	ΑE,	AG,	AL,	AM,	AT,	AU,	ΑZ,	BA,	BB,	BG,	BR,	BY,	BZ,	CA,	CH,	CN,
		co,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	ES,	FI,	GB,	GD,	GE,	GH,
		GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	KP,	KR,	KZ,	LC,	LK,	LR,
							MD,							-		•	

```
PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ,
             UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY,
             KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES,
             FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, SE, SI, SK, TR, BF,
             BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
     CA 2474695
                          A1
                                20030814
                                             CA 2003-2474695
                                                                     20030131
     AU 2003216139
                          A1
                                20030902
                                             AU 2003-216139
                                                                     20030131
    US 2003211560
                                             US 2003-356665
                          A1
                                20031113
                                                                     20030131
    US 7148030
                          B2
                                20061212
                          A1
     EP 1472238
                                20041103
                                             EP 2003-737580
                                                                    20030131
             AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK
     JP 2005530485
                          Т
                                20051013
                                             JP 2003-565985
                                                                    20030131
    US 2006183177
                          A1
                                             US 2006-346043
                                20060817
                                                                     20060202
     US 2006121546
                          A1
                                20060608
                                             US 2006-347054
                                                                    20060203
PRIORITY APPLN. INFO .:
                                             US 2002-353158P
                                                                 P 20020201
                                             US 2003-356665
                                                                 A1 20030131
                                             WO 2003-US2936
                                                                 W 20030131
```

AB A sensitive bioluminescent assay to detect proteases including caspases, trypsin and tryptase is provided. The method comprises contacting a sample suspected of having one or more caspases with a mixture comprising beetle luciferase and an aminomodified beetle aminoluciferin or a carboxyterminal protected derivative thereof, wherein the amino group of aminoluciferin or the derivative thereof is modified so as to covalently link a substrate for the caspase via a peptide bond to aminoluciferin or the carboxyterminal protected derivative thereof. If the sample comprises a caspase having a recognition site in the substrate, the substrate is cleaved at the peptide bond that links the substrate to aminoluciferin, yielding aminoluciferin, a substrate for the luciferase, in the mixture Luminescence is then detected. The method further comprises correlating luminescence with protease concentration or activity, i.e., increased luminescence correlates with increased protease concentration or activity. Also provided is a compound comprising aminoluciferin or a carboxyterminal protected derivative thereof covalently linked via a peptide bond to a protease recognition site such as a caspase recognition site, a trypsin recognition site, or a tryptase recognition site. A specific compound of the invention is a compound of formula I (R = peptide with an aspartic acid, lysine, or arginine C-terminus; R' = H, carboxy protecting group, e.g., C1-6-alkyl, Ph, benzyl ester, counterion). The invention also provides synthetic processes and intermediates disclosed herein, which are useful for preparing compds. of the invention. As described herein below, using a substrate for caspase 3 and 7 that was linked to either aminoluciferin or rhodamine-110, it was found that the limit of detection for the aminoluciferin-based substrate was 0.2 to 0.5 μU of purified caspase while that for the rhodamine-110-based substrate was 10 μU . As also described herein, it was found that the limit of detection of caspase expressing cells with the aminoluciferin-based substrate was 15 cells at 1 h while the limit of detection for the rhodamine-1 10-based substrate was 150 cells at 1 h.

REFERENCE COUNT: 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

```
L22 ANSWER 13 OF 24 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 13 ACCESSION NUMBER: 2003:376823 HCAPLUS Full-text
```

DOCUMENT NUMBER: 138:365147

TITLE: Compositions, methods and kits pertaining to

luminescent compounds

INVENTOR(S): Wood, Keith; Hawkins, Erika; Scurria, Mike; Klaubert, Dieter

PATENT ASSIGNEE(S): Promega Corporation, USA SOURCE: PCT Int. Appl., 60 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent English

LANGUAGE: FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

P		ENT 1				KIN		DATE				ICAT				D.	ATE	
w		2003							•							2	0021	101
												BG,						
			CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	ES,	FI,	GB,	GD,	GE,	GH,
			GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	KP,	KR,	KZ,	LC,	LK,	LR,
			LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NO,	NZ,	OM,	PH,
			PL,	PT,	RO,	RU,	SD,	SE,	SG,	SI,	SK,	SL,	ТJ,	TM,	TN,	TR,	TT,	TZ,
			UA,	ŬĠ,	UZ,	VC,	VN,	YU,	ZA,	ZM,	ZW							
		RW:	GH,	GM,	KE,	LS,	MW,	MZ,	SD,	SL,	SZ,	TZ,	UG,	ZM,	ZW,	ΑT,	BE,	BG,
			CH,	CY,	CZ,	DE,	DK,	EE,	ES,	FI,	FR,	GB,	GR,	IE,	IT,	LU,	MC,	NL,
			PT,	SE,	SK,	TR,	BF,	ВJ,	CF,	CG,	CI,	CM,	GA,	GN,	GQ,	GW,	ML,	MR,
				•	TD,													
U	S	2003:	1530	90		A1		2003	0814	1	US 2	001-	5348	2		2	0011	102
		24625						2003	0515	(CA 2	002-	2462	506		2	0021	101
		20023										002-						
Ē	P	1451:	155			A1		2004	0901	1	EP 2	002-	8028	15		2	0021	101
		R:	AT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR,	IT,	LI,	LU,	NL,	SE,	MC,	PT,
			IE,	SI,	LT,	LV,	FI,	RO,	MK,	CY,	AL,	TR,	BG,	CZ,	EE,	SK		
		16128	-									002-						
J	P	20055	5159	77		T	:	2005	0602		JP 2	003-	5421	46		2	0021	101
PRIORI	ΤY	APPI	LN.	INFO	.:					1	US 2	001-	53482	2	7	A 2	0011	102
										1	WO 2	002-	JS349	972	1	W 2	0021	101

OTHER SOURCE(S): MARPAT 138:365147

AB A method of measuring the enzymic activity of a luciferase includes contacting a luminogenic protein, such as a luciferase, with a protected luminophore to form a composition; and detecting light produced from the composition The protected luminophore provides increased stability and improved signal-to-background ratios relative to the corresponding unmodified coelenterazine.

REFERENCE COUNT: 8 THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L22 ANSWER 14 OF 24 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 14

ACCESSION NUMBER:

2001:924094 HCAPLUS Full-text

DOCUMENT NUMBER:

136:50649

TITLE:

Method for increasing luminescence assay sensitivity

INVENTOR(S): Hawkins, Erika; Centanni, John M.; Sankbeil,

Jacqueline; Wood, Keith V.

PATENT ASSIGNEE(S):

Promega Corporation, USA

SOURCE:

PCT Int. Appl., 45 pp. CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT NO.	KIND DATE	E APPLICAT	ION NO.	DATE
WO 2001096862	A2 2001	11220 WO 2001-	US18363	20010607
WO 2001096862	A3 2002	20718		
W: AE, AG, AL,	, AM, AT, AU,	, AZ, BA, BB, BG,	BR, BY, BZ,	CA, CH, CN,
CO, CR, CU	, CZ, DE, DK,	, DM, DZ, EE, ES,	FI, GB, GD,	GE, GH, GM,
HR, HU, ID	, IL, IN, IS,	, JP, KE, KG, KP,	KR, KZ, LC,	LK, LR, LS,
LT, LU, LV	, MA, MD, MG,	, MK, MN, MW, MX,	MZ, NO, NZ,	PL, PT, RO,

```
RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ,
             VN, YU, ZA, ZW
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
             DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF,
             BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
     US 7118878
                          B1
                                 20061010
                                             US 2000-590884
                                                                     20000609
     CA 2411179
                          A1
                                 20011220
                                             CA 2001-2411179
                                                                    20010607
                          A2
     EP 1297337
                                20030402
                                             EP 2001-942027
                                                                     20010607
             AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO, MK, CY, AL, TR
     JP 2004503777
                          Т
                                20040205
                                             JP 2002-510941
                                                                     20010607
     US 2004096924
                          A1
                                20040520
                                             US 2003-692587
                                                                    20031024
     US 7078181
                          B2
                                20060718
     US 2006051827
                          A1
                                20060309
                                             US 2004-991759
                                                                     20041118
     US 7108996
                          B2
                                20060919
PRIORITY APPLN. INFO.:
                                             US 2000-590884
                                                                 A 20000609
                                                                 W 20010607
                                             WO 2001-US18363
                                             US 2003-692587
                                                                 A3 20031024
```

AB A method for increasing the sensitivity of a luminescent assay comprising carrying out the assay in the presence of an organic compound that reduces luminescence that is not dependent on the presence of an analyte by at least about 10 fold, and that reduces luminescence that is dependent on the presence of an analyte by less than about 7 fold.

L22 ANSWER 15 OF 24 HCAPLUS COPYRIGHT 2007 ACS on STN ACCESSION NUMBER: 2003:108790 HCAPLUS Full-text

DOCUMENT NUMBER:

139:129758

TITLE:

Coelenterazine derivatives for improved solution

solubility

AUTHOR(S):

Hawkins, Erika M.; O'Grady, Michael;
Klaubert, Dieter; Scurria, Michael;

Good, Troy; Stratford, Cathy; Flemming, Rod; Simpson,

Dan; Wood, Keith V.

CORPORATE SOURCE:

SOURCE:

Promega Corporation, Madison, WI, 53715, USA

Bioluminescence & Chemiluminescence: Progress &

Current Applications, [Proceedings of the Symposium on

Bioluminescence and Chemiluminescence], 12th, Cambridge, United Kingdom, Apr. 5-9, 2002 (2002),

149-152. Editor(s): Stanley, Philip E.; Kricka, Larry J. World Scientific Publishing Co. Pte. Ltd.:

Singapore, Singapore.

CODEN: 69DPGZ; ISBN: 981-238-156-2

DOCUMENT TYPE: Conference LANGUAGE: English

AB Intracellular luminescent techniques requiring coelenterazine, such as bioluminescence resonance energy transfer (BRET), calcium detection, and intracellular reporter measurements, must accommodate the poor stability of this substrate in physiol. buffered solns. Coelenterazine degradation leads both to loss of luminescence over time, and increased background luminescence caused by enzyme-independent oxidation (autoluminescence). Both conditions limit luminescence sensitivity by reducing the signal-to-noise ratio. Coelenterazine can be stabilized by derivatizing the enol oxygen with an acyl oxymethyl ether. This prevents spontaneous oxidation in solution while making the substrate available intracellularly upon cleavage of the blocking group by endogenous esterases. We will describe the stability of pivaloyl oxymethyl coelenterazine-h (POM coelenterazine-h), and the effect of POM coelenterazine-h on intracellular luminescence, autoluminescence, and luminescent reaction kinetics. Also, we will present the characteristics of two other coelenterazine derivs.

10/053,482 April 30, 2007

L22 ANSWER 16 OF 24 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on

STN

ACCESSION NUMBER: 2007:78327 BIOSIS Full-text

DOCUMENT NUMBER: PREV200700078755

TITLE: Bioluminescent protease assay.

AUTHOR(S): Anonymous; O'Brien, Martha [Inventor]; Wood, Keith

V. [Inventor]; Klaubert, Dieter [Inventor];

Daily, William [Inventor]

CORPORATE SOURCE: Madison, WI USA

ASSIGNEE: Promega Corporation

PATENT INFORMATION: US 07148030 20061212

SOURCE: Official Gazette of the United States Patent and Trademark

Office Patents, (DEC 12 2006) CODEN: OGUPE7. ISSN: 0098-1133.

DOCUMENT TYPE: Patent LANGUAGE: English

ENTRY DATE: Entered STN: 24 Jan 2007

Last Updated on STN: 24 Jan 2007

AB A sensitive bioluminescent assay to detect proteases including caspases,

trypsin and tryptase is provided.

L22 ANSWER 17 OF 24 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on

STN

ACCESSION NUMBER: 2007:39141 BIOSIS Full-text

DOCUMENT NUMBER: PREV200700045138

TITLE: Method for increasing luminescence assay sensitivity.

AUTHOR(S): Anonymous; Hawkins, Erika [Inventor]; Centanni,

John M. [Inventor]; Sankbeil, Jacqueline [Inventor];

Wood, Keith V. [Inventor]

CORPORATE SOURCE: Madison, WI USA

ASSIGNEE: Promega Corporation

PATENT INFORMATION: US 07118878 20061010

SOURCE: Official Gazette of the United States Patent and Trademark

Office Patents, (OCT 10 2006) CODEN: OGUPE7. ISSN: 0098-1133.

DOCUMENT TYPE: Patent LANGUAGE: English

ENTRY DATE: Entered STN: 3 Jan 2007

Last Updated on STN: 3 Jan 2007

The invention provides kits and methods for increasing the sensitivity of a bio-luminescent assay, which employ an organic compound that, for instance, reduces luminescence that is not dependent on the presence of an analyte by at least about 10 fold and reduces luminescence that is dependent on the presence of an analyte by less than about 7 fold, reduces luminescence generated by luminogenic molecules not bound to an enzyme by at least about 10 fold and reduces the luminescence generated by luminogenic molecules bound to an enzyme by less than about 7 fold, or reduces autoluminescence by at least about 10 fold and reduces luminescence that is dependent on the presence of an analyte by less than about 7 fold.

L22 ANSWER 18 OF 24 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on

STN

ACCESSION NUMBER: 2007:23679 BIOSIS Full-text

DOCUMENT NUMBER: PREV200700032812

TITLE: Method for increasing luminescence assay sensitivity.

AUTHOR(S): Anonymous; Hawkins, Erika [Inventor]; Centanni,

John M. [Inventor]; Sankbeil, Jacqueline [Inventor];

Wood, Keith V. [Inventor]

CORPORATE SOURCE: Madison, WI USA

ASSIGNEE: Promega Corporation

PATENT INFORMATION: US 07108996 20060919

SOURCE: Official Gazette of the United States Patent and Trademark

> Office Patents, (SEP 19 2006) CODEN: OGUPE7. ISSN: 0098-1133.

DOCUMENT TYPE:

Patent

LANGUAGE:

English

ENTRY DATE:

Entered STN: 27 Dec 2006

Last Updated on STN: 27 Dec 2006

AB A method for increasing the sensitivity of a bio-luminescent assay comprising carrying out the assay in the presence of an organic compound that reduces luminescence that is not dependent on the presence of an analyte by at least about 10 fold, and that reduces, maintains, or increases the luminescence that is dependent on the presence of an analyte.

L22 ANSWER 19 OF 24 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on

STN

ACCESSION NUMBER:

2006:669792 BIOSIS Full-text

DOCUMENT NUMBER:

PREV200600682021

TITLE: AUTHOR(S): Kits for increasing luminescence assay sensitivity. Anonymous; Hawkins, Erika [Inventor]; Centanni,

John M. [Inventor]; Sankbeil, Jacqueline [Inventor];

Wood, Keith V. [Inventor]

CORPORATE SOURCE:

Madison, WI USA

ASSIGNEE: Promega Corporation

PATENT INFORMATION: US 07078181 20060718

SOURCE:

Official Gazette of the United States Patent and Trademark

Office Patents, (JUL 18 2006) CODEN: OGUPE7. ISSN: 0098-1133.

DOCUMENT TYPE:

Patent

LANGUAGE:

English

ENTRY DATE:

Entered STN: 6 Dec 2006

Last Updated on STN: 6 Dec 2006

Assay kits for increasing the sensitivity of luminescent assays are provided AB that include an organic compound that reduces luminescence that is not dependent on the presence of an analyte by at least about 10 fold, and that reduces, maintains, or increases the luminescence that is dependent on the presence of an analyte.

L22 ANSWER 20 OF 24 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on

STN

ACCESSION NUMBER:

2006:584734 BIOSIS Full-text

DOCUMENT NUMBER:

PREV200600595360

TITLE: AUTHOR(S): Homogeneous, bioluminescent assays for proteasome activity. O'Brien, Martha A. [Reprint Author]; Moravec, Richard A.;

Riss, Terry; Scurria, Michael A.; Daily, William

J.; Klaubert, Dieter H.; Wood, Keith V.

; Bulleit, Robert F.

CORPORATE SOURCE:

Promega Corp, Madison, WI USA

SOURCE:

Proceedings of the American Association for Cancer Research

Annual Meeting, (APR 2006) Vol. 47, pp. 321. Meeting Info.: 97th Annual Meeting of the

American-Association-for-Cancer-Research (AACR). Washington, DC, USA. April 01 -05, 2006. Amer Assoc Canc

Res.

10/053,482 April 30, 2007

ISSN: 0197-016X.

DOCUMENT TYPE: Conference; (Meeting)

Conference; Abstract; (Meeting Abstract)

LANGUAGE: English

ENTRY DATE: Entered STN: 8 Nov 2006

Last Updated on STN: 8 Nov 2006

L22 ANSWER 21 OF 24 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on

STN

ACCESSION NUMBER: 2007:261852 BIOSIS Full-text

DOCUMENT NUMBER: PREV200700271919

TITLE: A multipurpose fusion protein tag for analysis of dynamic

cellular events.

AUTHOR(S): Learish, Randall D. [Reprint Author]; Los, Georgyi V.;

Zimprich, Chad; McDougall, Mark; Karassina, Natasha; Nath,

Nidhi; Darzins, Al; Klaubert, Dieter; Bulleit,

Robert F.; Wood, Keith

CORPORATE SOURCE: Promega Corp, Madison, WI USA

SOURCE: Proceedings of the American Association for Cancer Research

Annual Meeting, (APR 2005) Vol. 46, No. Suppl. S, pp. 902.

Meeting Info.: 96th Annual Meeting of the

American-Association-for-Cancer-Research. Anaheim, CA, USA.

April 16 -20, 2005. Amer Assoc Canc Res.

ISSN: 0197-016X.

DOCUMENT TYPE: Conference; (Meeting)

Conference; Abstract; (Meeting Abstract)

LANGUAGE: English

ENTRY DATE: Entered STN: 25 Apr 2007

Last Updated on STN: 25 Apr 2007

L22 ANSWER 22 OF 24 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on

STN

ACCESSION NUMBER: 2006:219506 BIOSIS Full-text

DOCUMENT NUMBER: PREV200600222633

TITLE: The HaloTag (TM): a novel technology for cellular analysis.

AUTHOR(S): Los, G. V. [Reprint Author]; Zimprich, C.; McDougall, M.

G.; Karassina, N.; Learish, R.; Klaubert, D. H.;

Darzins, A.; Bulleit, R. F.; Wood, K.

CORPORATE SOURCE: Promega Corp, Madison, WI USA

SOURCE: Journal of Neurochemistry, (JUN 2005) Vol. 94, No. Suppl.

1, pp. 15.

Meeting Info.: 36th Annual Meeting of the

American-Society-for-Neurochemistry. Madison, WI, USA. June

25 -29, 2005. Amer Soc Neurochem. CODEN: JONRA9. ISSN: 0022-3042.

DOCUMENT TYPE: Conference; (Meeting)

Conference; Abstract; (Meeting Abstract)

LANGUAGE: English

ENTRY DATE: Entered STN: 5 Apr 2006

Last Updated on STN: 5 Apr 2006

L22 ANSWER 23 OF 24 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on

STN

ACCESSION NUMBER: 2004:195934 BIOSIS Full-text

DOCUMENT NUMBER: PREV200400196493

TITLE: Site - specific localization of small molecule reporters

within living cells.

AUTHOR(S): Los, G. V. [Reprint Author]; Zimprich, C. [Reprint Author];

McDougall, M. G.; Karassina, N. [Reprint Author]; Learish,

R. [Reprint Author]; Klaubert, D.; Wood,

10/053,482 April 30, 2007

K. [Reprint Author]; Bulleit, B. [Reprint Author]

CORPORATE SOURCE: R&D, Promega Corp., Madison, WI, USA

SOURCE: Society for Neuroscience Abstract Viewer and Itinerary

Planner, (2003) Vol. 2003, pp. Abstract No. 232.5.

http://sfn.scholarone.com. e-file.

Meeting Info.: 33rd Annual Meeting of the Society of Neuroscience. New Orleans, LA, USA. November 08-12, 2003.

Society of Neuroscience.

DOCUMENT TYPE: Conference; (Meeting)

Conference; Abstract; (Meeting Abstract)

LANGUAGE: English

ENTRY DATE: Entered STN: 14 Apr 2004

Last Updated on STN: 14 Apr 2004

AB The ability to specifically label proteins can help reveal information about protein functions and dynamics within the complex biochemical environment of the living cells. Here we describe a novel technology for covalently tethering functional groups ((FG), e.g. fluorescent dye) to a universal reporting protein (URP) within cells. The URP is a mutant enzyme capable of forming a stable covalent bond to a modified substrate coupled to the functional group. In our initial approach the URP is a halo-alkane dehalogenase from Rhodococcus rhodochrous (DhaA) with a mutation in a critical active site residue. The activity of DhaA cleaves carbon-halogen bonds in aliphatic and aromatic halogenated compounds involving a typical hydrolytic triad. In this reaction an enzyme-substrate complex is formed by a nucleophilic attack involving Asp106 and the formation of an ester intermediate; His272 activates H2O that hydrolyzes this intermediate releasing product from the catalytic center. A point mutation in DhaA involving a substitution of phenylalanine for His272 impairs the hydrolysis step leading to a stable covalent intermediate with substrate and any conjugated FG. This technology allows the labeling of mutant DhaA or mutant DhaA fusion proteins expressed in cells. A significant advantage of this approach is the flexibility to create labeled proteins with a potentially wide range of optical properties or other functionalities. The technology can be applied in different cells and organisms allowing a variety of experimental approaches to study protein function in living cells.

L22 ANSWER 24 OF 24 WPIX COPYRIGHT 2007 THE THOMSON CORP on STN

ACCESSION NUMBER: 2006-135434 [14] WPIX CROSS REFERENCE: 2004-635192; 2006-044483

DOC. NO. CPI: C2006-046443 [14]

TITLE: New mutant dehalogenase comprising at least two amino

acid substitutions relative to a corresponding wild-type

dehalogenase, useful for isolating, detecting,

identifying, imaging, displaying, or localizing molecules

of interest

DERWENT CLASS: B04; D16

INVENTOR: DARZINS A; ENCELL L; KLAUBERT D; LOS GEORGYI V;

MCDOUGALL M; WOOD K V; WOOD M G; ZIMPRICH C

PATENT ASSIGNEE: (DARZ-I) DARZINS A; (ENCE-I) ENCELL L; (KLAU-I) KLAUBERT

D; (LGEO-I) LOS GEORGYI V; (MCDO-I) MCDOUGALL M; (WOOD-I)

WOOD K V; (WOOD-I) WOOD M G; (ZIMP-I) ZIMPRICH C

COUNTRY COUNT: 1

PATENT INFO ABBR.:

PATENT NO KIND DATE WEEK LA PG MAIN IPC

US 20060024808 A1 20060202 (200614)* EN 170[60]

APPLICATION DETAILS:

PAT	TENT NO	KI	ND 	API	PLICATION	DATE
US	20060024	1808 A1	Provisional	US	2004-5924991	20040730
US	20060024	1808 A1		US	2005-194110	20050729

PRIORITY APPLN. INFO: US 2005-194110 20050729 US 2004-592499P 20040730

AB US 20060024808 A1 UPAB: 20060227

NOVELTY - A mutant dehalogenase comprising at least two amino acid substitutions relative to a corresponding wild-type dehalogenase, where the mutant dehalogenase forms a bond with a dehalogenase substrate which comprises one or more functional groups, which bond is more stable than the bond formed between the corresponding wild-type dehalogenase and the substrate, is new.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for:

- (1) a method for detecting or determining the presence or amount of a mutant hydrolase;
- (2) a method for isolating a molecule, cell or sub-cellular organelle of interest in a sample;
 - (3) a method for labeling a cell;
 - (4) a method for labeling a cell;
 - (5) a polynucleotide encoding the mutant hydrolase;
- (6) a mutant hydrolase comprising at least two amino acid substitutions relative to a corresponding wild-type hydrolase, where one substitution is at a position corresponding to amino acid residue 272 of a Rhodococcus rhodochrous dehalogenase or at a position corresponding to amino acid residue 106 of a Rhodococcus rhodochrous dehalogenase, and a second substitution is at an amino acid residue corresponding to position 175, 176 or 273 of a Rhodococcus rhodochrous dehalogenase; and
- (7) a thermostable mutant dehalogenase comprising at least one substitution at a position corresponding to amino acid residue 175 of a Rhodococcus rhodochrous dehalogenase, which substitution is correlated with enhanced thermostability relative to a corresponding mutant dehalogenase without the substitution at the position corresponding to amino acid residue 175; and a compound of formula XXIX-XXXIV.
- USE The mutant dehalogenase is useful for isolating, detecting, identifying, imaging, displaying, or localizing molecules of interest; labeling cells, including live cell imaging; or labeling proteins in vitro and/or in vivo.

SEARCH HISTORY

```
=> d his nofil
```

(FILE 'HOME' ENTERED AT 10:29:26 ON 30 APR 2007)

FILE 'HCAPLUS' ENTERED AT 10:29:29 ON 30 APR 2007

E US2001-053482/APPS

1 SEA ABB=ON PLU=ON (US2001-53482/AP OR US2001-53482/PRN) SEL RN

FILE 'REGISTRY' ENTERED AT 10:30:06 ON 30 APR 2007

L2 11 SEA ABB=ON PLU=ON (50909-86-9/BI OR 524066-91-9/BI OR 524066-92-0/BI OR 524066-93-1/BI OR 524066-94-2/BI OR 524066-95 -3/BI OR 524066-96-4/BI OR 55779-48-1/BI OR 61869-41-8/BI OR 65417-16-5/BI OR 70217-82-2/BI)

FILE 'REGISTRY' ENTERED AT 10:30:41 ON 30 APR 2007

L3 STR

L4 0 SEA SSS SAM L3

L5 8 SEA SSS FUL L3

FILE 'HCAPLUS' ENTERED AT 10:33:01 ON 30 APR 2007

L6 4 SEA ABB=ON PLU=ON L5

FILE 'BEILSTEIN' ENTERED AT 10:33:14 ON 30 APR 2007

L7 0 SEA SSS SAM L3

L8 1 SEA SSS FUL L3

L9 1 SEA ABB=ON PLU=ON L8 AND RN/FA

L10 0 SEA ABB=ON PLU=ON L8 AND BABSAN/FA

SEL RN L8 D COST

FILE 'REGISTRY' ENTERED AT 10:34:25 ON 30 APR 2007

L11 1 SEA ABB=ON PLU=ON 65417-16-5/RNX

L12 1 SEA ABB=ON PLU=ON L11 AND L2

L13 1 SEA ABB=ON PLU=ON L11 AND L5

FILE 'MARPAT' ENTERED AT 10:35:26 ON 30 APR 2007

L14 0 SEA SSS SAM L3

L15 2 SEA SSS FUL L3

L16 1 SEA ABB=ON PLU=ON L15 NOT L6

FILE 'HCAPLUS, MEDLINE, EMBASE, BIOSIS, DISSABS, WPIX' ENTERED AT 10:36:23 ON 30 APR 2007

L17 2610 SEA ABB=ON PLU=ON WOOD K/AU OR WOOD K ?/AU OR WOOD KEITH?/AU

L*** DEL 938 S HAWKINS E/AU OR HAWKINS E ?/AU OR HAWKINS ERI!A/AU

L18 948 SEA ABB=ON PLU=ON HAWKINS E/AU OR HAWKINS E ?/AU OR HAWKINS ERI!A?/AU

E SCURRIA M/AU

L19 34 SEA ABB=ON PLU=ON ("SCURRIA M"/AU OR "SCURRIA M A"/AU OR "SCURRIA M S"/AU OR "SCURRIA MICHAEL"/AU OR "SCURRIA MICHAEL A"/AU OR "SCURRIA MIKE"/AU)

E KLAUBERT D/AU

L20 198 SEA ABB=ON PLU=ON ("KLAUBERT D"/AU OR "KLAUBERT D H"/AU OR "KLAUBERT D K"/AU OR "KLAUBERT DIETER H"/AU OR "KLAUBERT DIETER HEINZ"/AU)

L21 44 SEA ABB=ON PLU=ON (L17 AND (L18 OR L19 OR L20)) OR (L18 AND

(L19 OR L20)) OR (L19 AND L20)

FILE 'HCAPLUS' ENTERED AT 10:45:16 ON 30 APR 2007

D QUE L6

D L6 IBIB ABS HITSTR TOT

FILE 'MARPAT' ENTERED AT 10:45:32 ON 30 APR 2007

D QUE L16

D L16 IBIB ABS QHIT TOT

FILE 'HCAPLUS, MEDLINE, EMBASE, BIOSIS, DISSABS, WPIX' ENTERED AT 10:46:23 ON 30 APR 2007

D QUE

L22

24 DUP REM L21 (20 DUPLICATES REMOVED)

ANSWERS '1-15' FROM FILE HCAPLUS ANSWERS '16-23' FROM FILE BIOSIS

ANSWER '24' FROM FILE WPIX

D L22 IBIB AB TOT